

SUPPLEMENTAL TABLE 1

Two independent duplicate T_{50} values ($^{\circ}\text{C}$) for parent CBH IIs, 23 original sample set CBH II chimeras and predicted thermostable CBH II chimeras. The 18 chimeras synthesized for this work are preceded by an asterisk.

| Sample Set Chimeras & Parents | | | | Predicted Thermostable Chimeras | | | |
|-------------------------------|-------------|-------------|---------------|---------------------------------|-------------|-------------|---------------|
| Sequence | $T_{50}(1)$ | $T_{50}(2)$ | Mean T_{50} | Sequence | $T_{50}(1)$ | $T_{50}(2)$ | Mean T_{50} |
| 32333113 | 52 | 51 | 51.5 | 12332331 | 66.5 | 67 | 66.8 |
| 13111313 | 56 | 53.5 | 54.8 | *13112332 | 67 | 67 | 67 |
| 11313121 | 55 | 55.5 | 55.3 | 22311331 | 68 | 68 | 68 |
| 21131311 | 57.5 | 57 | 57.3 | *12111332 | 68 | 68 | 68 |
| 31212111 | 59 | 58 | 58.5 | *12112332 | 68.5 | 67.5 | 68 |
| Parent 2 | 60 | 58 | 59 | 12131331 | 68.5 | 69 | 68.8 |
| 23233133 | 61 | 61 | 61 | *12131332 | 70 | 67.5 | 68.8 |
| 31311112 | 60 | 62 | 61 | *12332332 | 69 | 69 | 69 |
| 22212231 | 63 | 61 | 62 | 12111131 | 70 | 68.5 | 69.3 |
| 13231111 | 63 | 6.5 | 63.3 | 12311332 | 70 | 69 | 69.5 |
| 12213111 | 63 | 63.5 | 63.3 | 13332331 | 70 | 69 | 69.5 |
| Parent 3 | 63.5 | 64.5 | 64 | 12132331 | 70.5 | 69 | 69.8 |
| 12133333 | 64 | 64 | 64 | *12132332 | 70.5 | 69 | 69.8 |
| Parent 1 | 64 | 65.5 | 64.8 | *13332332 | 69.5 | 70 | 69.8 |
| 33133132 | 65 | 66 | 65 | 12112132 | 71 | 68.5 | 69.8 |
| 11332333 | 64.5 | 66 | 65.3 | 13322332 | 71 | 68.5 | 69.8 |
| 23311333 | 65 | 66 | 65.5 | *13131332 | 70 | 70 | 70 |
| 33213332 | 66 | 66 | 66 | *12331332 | 71 | 69 | 70 |
| 13333232 | 67.5 | 67 | 67.3 | *13312332 | 70 | 70 | 70 |
| 22232132 | 68 | 68 | 68 | *11113332 | 69.5 | 70.5 | 70 |
| 11113132 | 71.5 | 71 | 71.3 | *13113132 | 70.5 | 69.5 | 70 |
| 21333331 | 73.5 | 75.5 | 74.5 | *11112132 | 70.5 | 70 | 70.3 |
| 21311131 | 75.5 | 75.5 | 75 | *12113132 | 70.5 | 70.5 | 70.5 |
| | | | | *13132332 | 69.5 | 71.5 | 70.5 |
| | | | | *11111132 | 71 | 70.5 | 70.8 |
| | | | | 13331332 | 72 | 70 | 71 |
| | | | | *13111132 | 72 | 69.5 | 71.3 |
| | | | | *12222132 | 72.5 | 70 | 71.3 |
| | | | | 12222332 | 72 | 69.5 | 71.3 |
| | | | | 13311332 | 71 | 71.5 | 71.7 |
| | | | | 13311331 | 73.5 | 72.5 | 73 |

SUPPLEMENTAL TABLE 2

Amino acid sequences for CBH II parent and chimera catalytic domains shown in Supplemental Table 1. Table also includes catalytic domain for *P. chrysosporium* CBH II as described in text. All recombinant CBH IIs share the N-terminal CBM and linker from the native *H. jecorina* CBH II, CSSVWGQCGGNWSGPTCCASGSTCVYSNDYYSQCLPGAASSSSSTRAASTTSRVSPSTTSRSSSA TPPPGSTTTRVPPVSGGTATYS.

Parent 1 (*H. insolens*)

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFQWLDRNVTVDTLVQTLSE
IREANQAGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIREILISFSDVRT
ILVIEPDSL ANMVTNMNV PKCSGAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGWPANI Q
PAAELFAKIYEDAGKPRAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKH YIEAFRPLLEARGF
P_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDGTS
DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

Parent 2 (*H. jecorina*)

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFMWLD T_LDKTPLMEQTLA
DIRTANKNGGN_YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDTIRQIVVEYS DIR
TLLVIEPDSL ANLVTNLGTPKCANAQSAYLE CINYAVTQLNLPNVAMYLDAGHAGWLGWPAN Q
DPAAQLFANVYKNASSPRALRGLATNVANYNGWNITSPPSYTQGN AVYNEKLYIHAIGPLL ANH
GWSNAFFITDQGRSGKQPTGQQQWGDWCNVIGTGFGIRPSANTGDSLLDSFVWVKPGGECDGT
SDSSAPRFD SHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

Parent 3 (*C. thermophilum*)

GNPFSGVQLWANTYYSSEVHTLAIPSL S_PELAAKAAKVAEVPSFQWLDRNVTVDTLFSGTLAEI
RAANQRGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIRELLIQYSDIRTI
LVIEPDSL ANMVTNMNVQKCSNAASTYKELTVYALKQLNLPNVAMYMDAGHAGWLGWPANI
QPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDTSAARYDYHCGLS DALTPAPEAGQWFQAYFEQLLINANPPF_

P. chrysosporium CBH II

NNPWTGFQIFLSPYYANEVAAAA_KQITDPTLSSKAASVAN IPTFTWLDSVAKIPDLGTYLASASA
LGKSTGT_KQLVQIVYDLPDRDCAAKASNGEFSIANNGQANYENYIDQIVAQIQQFPDVRVVA
VIEPDSL ANLVTNLNVQKCANAKTTYLACVNYALTNLAKVGVYMYMDAGHAGWLGWPANLS
PAAQLFTQVWQNAGKSPFIKGLATNVANYNALQAASPD PITQGNPNYDEIHYINALAPLLQQAG
WD_ATFIVDQGRSGVQNI RQ_QWGDWCNIKGAGFGTRPTTNTGSQFIDSIVWVKPGGECDGTSN
SSSPRYDSTCSLPDAAQPAPEAGTWFQAYFQTLVSAANPPL_

32333113

GNPFSGVQLWANTYYSSEVHTLAIPSL S_PELAAKAAKVAEVPSFMWLD T_LDKTPLMEQTLADI
RTANKNGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIRELLIQYSDIRTI
LVIEPDSL ANMVTNMNVQKCSNAASTYKELTVYALKQLNLPNVAMYMDAGHAGWLGWPANI
QPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKH YIEAFRPLLEAR
GFP_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDG
TSDTSAARYDYHCGLS DALTPAPEAGQWFQAYFEQLLINANPPF_

13111313

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFQWLDRNVTVDTLFSGTLAE

SCHEMA recombination uncovers stabilizing cellulase mutation

IRAANQRGANPQYAAQIVVYDLPDRDCAAAAASNGEWAIANNGVNNYKAYINRIREILISFSDVRT
ILVIEPDSLANMVTNMNVKPCSGAASTYRELTIIYALKQLDLPHVAMYMDAGHAGWLGWPANIQ
PAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKHIEAFAPLLRNQGF
D_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDGTS
DTSAARYDYHCGLSDALTPAPEAGQWFQAYFEQLLINANPPF_

11313121

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVAEPVSFQWLDRNVTVDTLVQTLSE
IREANQAGANPPYAGIFVVYDLPDRDCAAAAASNGEWSIANNGANNYKRYIDRIREILISFSDVRTI
LVIEPDSLANMVTNMNVKPCSGAASTYKELTVYALKQLNLPVAMYMDAGHAGWLGWPANI
QPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEAR
GFPNAFFITDQGRSGKQPTGQQQWGDWCNVIGTGFGIRPSANTGDSLLDSFVWVKPGGECDGTS
DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

21131311

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFQWLDRNVTVDTLVQTLSEI
EIREANQAGANPQYAAQIVVYDLPDRDCAAAAASNGEWAIANNGVNNYKAYINRIRELLIQYSDI
RTILVIEPDSLANMVTNMNVQKCSNAASTYRELTIIYALKQLDLPHVAMYMDAGHAGWLGWPA
NIQPAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKHIEAFAPLLRN
QGFD_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECD
GTSDDTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

31212111

GNPFSGVQLWANTYYSEVHTLAIPSLT_PELAAKAAKVAEVPSFQWLDRNVTVDTLVQTLSEI
REANQAGGN__YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIREILISFSDVRTIL
VIEPDSLANMVTNMNVKPCSGAASAYLECINYAVTQLNLPNVAMYLDAGHAGWLGWPANQDP
AAQLFANVYKNASSPRALRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEARGF
P_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDGTS
DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

23233133

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFQWLDRNVTVDTLFSGTLA
EIRAANQRRGN__YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIRELLIQYSDIR
TILVIEPDSLANMVTNMNVQKCSNAASTYKELTVYALKQLNLPVAMYMDAGHAGWLGWPA
NIQPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEAR
RGFP_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVDADFVWVKPGGESD
GTSDDTAARYDYHCGLSDALTPAPEAGQWFQAYFEQLLINANPPF

31311112

GNPFSGVQLWANTYYSEVHTLAIPSLT_PELAAKAAKVAEVPSFQWLDRNVTVDTLVQTLSEI
REANQAGANPPYAGIFVVYDLPDRDCAAAAASNGEWSIANNGANNYKRYIDRIREILISFSDVRTIL
VIEPDSLANMVTNMNVKPCSGAASTYRELTIIYALKQLDLPHVAMYMDAGHAGWLGWPANIQP
AAELFAKIYEDAGKPRAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEARGFP
_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDGTSD
SSAPRFDShCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

22212231

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFMWLDL_LDKTPLMEQTLA
DIRTANKNGGN__YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIREILISFSDVR
TILVIEPDSLANMVTNMNVKPCSGAASAYLECINYAVTQLNLPNVAMYLDAGHAGWLGWPAN

SCHEMA recombination uncovers stabilizing cellulase mutation

QDPAAQLFANVYKNASSPRALRGLATNVANYNGWNITSPPSYTQGNNAVYNEKLYIHAIGPLLAN
HGWS_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVDADFVWVKPGGESD
GTSDDTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

13231111

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVAEPVSFQWLDRNVTVDTLFSGTLAE
IRAANQRGGN_YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIRELLIQYSDIRTI
LVIEPDSLANMVTNMNVQKCSNAASTYRELTIIYALKQLDLPHVAMYMDAGHAGWLGWPA
PAAELFAKIYEDAGKPRAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEARGF
P_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDGTS
DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

12213111

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVAEPVSFMWLDL_LDKTPLMEQTLAD
IRTANKNGGN_YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIREILISFSDVRTI
LVIEPDSLANMVTNMNVPKCSGAASTYKELTVYALKQLNLPVAMYMDAGHAGWLGWPA
QPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEAR
GFP_AQFIVDQGRSGKQPTGQKEWGHWCNAIGTGFGMRPTANTGHQYVDAFVWVKPGGECDG
TSDTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

12133333

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVAEPVSFMWLDL_LDKTPLMEQTLAD
IRTANKNGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVNNYKAYINRIRELLIQYSDIR
TILVIEPDSLANMVTNMNVQKCSNAASTYKELTVYALKQLNLPVAMYMDAGHAGWLGWPA
NIQPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKHIEAFAPLLRN
QGFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVDADFVWVKPGGESD
GTSDDTAARYDYHCGLSDALTPAPEAGQWFQAYFEQLLINANPPF_

33133132

GNPFSGVQLWANTYYSEVHTLAIPSLP_PELAAKAAKVAEPVSFQWLDRNVTVDTLFSGTLAEI
RAANQRGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVNNYKAYINRIRELLIQYSDIR
ILVIEPDSLANMVTNMNVQKCSNAASTYKELTVYALKQLNLPVAMYMDAGHAGWLGWPA
QPAAELFAQIYRDAGRPAAVRGLATNVANYNAWSVSSPPPYTSPNPNYDEKHIEAFRPLLEAR
GFP_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVDADFVWVKPGGESD
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

11332333

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVAEPVSFQWLDRNVTVDTLVQTLSE
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LVIEPDSLANMVTNMNVQKCSNAASAYLECINYAVTQLNLPNVAMYLDAGHAGWLGWPA
DPAAQLFANVYKNASSPRALRGLATNVANYNAWSIASPPSYTSPNPNYDEKHIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVDADFVWVKPGGESD
TSDTSAARYDYHCGLSDALTPAPEAGQWFQAYFEQLLINANPPF_

23311333

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFQWLDRNVTVDTLFSGTLA
EIRAANQRGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIREILISFSDVR
TILVIEPDSLANMVTNMNVPKCSGAASTYRELTIIYALKQLDLPHVAMYMDAGHAGWLGWPA
QPAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKHIEAFAPLLRNQ

GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDTSAARYDYHCGLSDALTPAPEAGQWFQAYFEQLLINANPPF_

33213332

GNPFSGVQLWANTYYSEVHTLAIPSLS_PELAAKAAKVAEVPSFQWLDRNVTVDTLFSGTLAEI
RAANQRGGN__YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIREILISFSDVRTI
LVIEPDSL ANMVTNMNVPKCSGAASTYKELTVYALKQLNLPHVAMYMDAGHAGWLGW PANI
QPAAELFAQIYRDAGRPA A VRGLATNV ANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

13333232

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFQWLDRNVTVDTLFSGTLAE
IRAANQRGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIRELLIQYSDIRTI
LVIEPDSL ANMVTNMNVQKCSNAASTYKELTVYALKQLNLPHVAMYMDAGHAGWLGW PANI
QPAAELFAQIYRDAGRPA A VRGLATNV ANYNGWNITSPPSYTQGN AVYNEKLYIHAIGPLL ANH
GWS_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

22232132

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFMWLD T_LDKTPLMEQTLA
DIRTANKNGGN__YAGQFVVYDLPDRDCAALASNGEYSIADGGVAKYKNYIDRIRELLIQYSDIR
TILVIEPDSL ANMVTNMNVQKCSNAASAYLECIN YAVTQLNLPNVAMYLDAGHAGWLGW PAN
QDPAAQLFANVYKNASSPRALRGLATNV ANYNAWSVSSPPPYTSPNPNYDEKH YIEAFRPLLEA
RGFP_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESD
GTS DSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

11113132

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFQWLDRNVTVDTL LVQTLSE
IREANQAGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIREILISFSDVRT
ILVIEPDSL ANMVTNMNVPKCSGAASTYKELTVYALKQLNLPHVAMYMDAGHAGWLGW PANI
QPAAELFAQIYRDAGRPA A VRGLATNV ANYNAWSVSSPPPYTSPNPNYDEKH YIEAFRPLLEAR
GFP_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

21333331

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFQWLDRNVTVDTL LVQTL S
EIREANQAGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIRELLIQYSDIR
TILVIEPDSL ANMVTNMNVQKCSNAASTYKELTVYALKQLNLPHVAMYMDAGHAGWLGW PA
NIQPAAELFAQIYRDAGRPA A VRGLATNV ANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRN
QGFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESD
GTS DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

21311131

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFQWLDRNVTVDTL LVQTL S
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ILVIEPDSL ANMVTNMNVPKCSGAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGW PANIQ
PAAELFAKIYEDAGKPRAVRGLATNV ANYNAWSVSSPPPYTSPNPNYDEKH YIEAFRPLLEARGF
P_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDGTS
DTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

12332331

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDT_LDKTPLMEQTLAD
IRTANKNGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIRELLIQYSDIRTI
LVIEPDSL ANMVTNMNVQKCSNAASAYLECIN YAVTQLNLPNVAMYLDAGHAGWLGW PANQ
DPAAQLFANVYKNASSPRALRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDG
TSDTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

13112332

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFQWLDRNVTVDTLFSGTLAE
IRAANQRGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIREILISFSDVRT
ILVIEPDSL ANMVTNMNVPKCSGAASAYLECIN YAVTQLNLPNVAMYLDAGHAGWLGW PANQ
DPAAQLFANVYKNASSPRALRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

22311331

GNPFVGVTPWANAYYASEVSSLAIPSLT_GAMATAAAAVAKVPSFMWLDT_LDKTPLMEQTLA
DIRTANKNGANPPYAGIFVVYDLPDRDCAAAASNGEWSIANNGANNYKRYIDRIREILISFSDVR
TILVIEPDSL ANMVTNMNVPKCSGAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGW PANI
QPAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDG
TSDTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

12111332

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDT_LDKTPLMEQTLAD
IRTANKNGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIREILISFSDVRT
ILVIEPDSL ANMVTNMNVPKCSGAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGW PANIQ
PAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQGF
D_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDGT
DSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

12112332

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDT_LDKTPLMEQTLAD
IRTANKNGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIREILISFSDVRT
ILVIEPDSL ANMVTNMNVPKCSGAASAYLECIN YAVTQLNLPNVAMYLDAGHAGWLGW PANQ
DPAAQLFANVYKNASSPRALRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

12131331

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDT_LDKTPLMEQTLAD
IRTANKNGANPQYAAQIVVYDLPDRDCAAAASNGEWAIANNGVN NYKAYINRIRELLIQYSDIR
TILVIEPDSL ANMVTNMNVQKCSNAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGW PANI
QPAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGH ELVDAFVWVKPGGESDG
TSDTTAARYDYHCGLEDALKPAPEAGQWFNEYFIQLLRNANPPF_

SCHEMA recombination uncovers stabilizing cellulase mutation

12131332

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDLDT_LDKTPLMEQTLAD
IRTANKNGANPQYAAQIVVYDLPDRDCAAAAASNGEWAIANNGVN NYKAYINRIRELLIQYSDIR
TILVIEPDSL ANMVTNMNVQKCSNAASTYRELTIYALKQLDLPHVAMYMDAGHAGWLGW PANI
QPAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKH YIEAFAPLLRNQ
GFD_AKFIVDTGRNGKQPTGQLEWGHWCNVKGTGFGVRPTANTGHELVD AFVWVKPGGESDG
TSDSSAPRFDSHCALPDALQPAPQAGAWFQAYFVQLLTNANPSFL

12332332

GNPFEGVQLWANNYYRSEVHTLAIPQITDPALRAAASAVA EVPSFMWLDLDT_LDKTPLMEQTLAD
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12111131

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12311332

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13332331

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12132331

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12132332

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SCHEMA recombination uncovers stabilizing cellulase mutation

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13332332

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12112132

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13322332

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13312332

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LVIEPDSLANMVTNMNVKCSGAASAYLECINYAVTQLNLPNVAMYLDAGHAGWLGWSPANQD

SCHEMA recombination uncovers stabilizing cellulase mutation

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11113332

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ILVIEPDSLANMVTNMNVKCSGAASTYKELTVYALKQLNLPHVAMYMDAGHAGWLGWSPANI
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13132332

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12222332

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13311332

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13311331

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LVIEPDSL ANMVTNMNVPKCSGAASTYRELT IYALKQLDLPHVAMYMDAGHAGWLGW PANIQ
PAAELFAKIYEDAGKPRAVRGLATNVANYNAWSIASPPSYTSPNPNYDEKHYIEAFAPLLRNQGF
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SUPPLEMENTAL TABLE 3

T₅₀ linear regression model parameters and p-values. Parameter values with $p \leq 0.1$, used to calculate the regression fit line of Figure 1, appear in bold. Block effects are reported relative to a parent 1 (*H. insolens* CBH II) reference state with 16 parameters representing substitution of each of the 8 blocks from parents 2 and 3.

| Block | Parameter Value | p-value |
|----------------|-----------------|-------------|
| Parent1 | 62.8 | 0.00 |
| B12 | -0.9 | 0.35 |
| B13 | -3.5 | 0.00 |
| B22 | -1.7 | 0.06 |
| B23 | -1.1 | 0.25 |
| B32 | 0.5 | 0.68 |
| B33 | 1.2 | 0.10 |
| B42 | 2.7 | 0.05 |
| B43 | 0.0 | 0.99 |
| B52 | -1.3 | 0.10 |
| B53 | -0.6 | 0.50 |
| B62 | -3.5 | 0.02 |
| B63 | -0.7 | 0.37 |
| B72 | -3.8 | 0.05 |
| B73 | 8.5 | 0.00 |
| B82 | 0.0 | 1.00 |
| B83 | -5.6 | 0.00 |

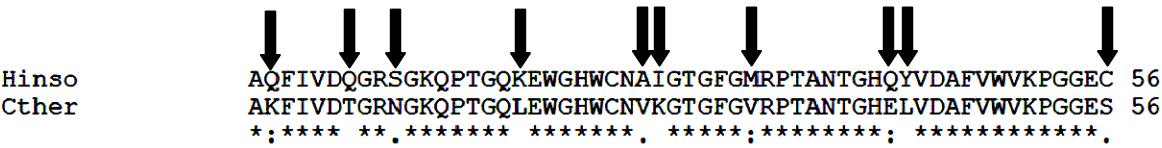
SUPPLEMENTAL TABLE 4

Total activity in synthetic (SDCAA) and rich (YPD) expression culture media supernatants for *H. jecorina* and *H. insolens* wild type, Ser point mutant and B7P3 block substitution CBH IIs. Values presented are μg glucose/mL cellulase activity assay per mL of expression culture supernatant CBH II equivalent added to cellulase activity assay. For SDCAA cultures, concentrated SDCAA culture supernatants were used and activity toward phosphoric acid swollen cellulose (1 mg/mL) at 50 °C for 100 minutes in 50 mM sodium acetate, pH 4.8, was measured. YPD supernatant CBH II was concentrated by binding to Avicel and activity toward Avicel (15 mg/mL) at 55 °C for 150 minutes in 50 mM sodium acetate, pH 4.8, was measured.

| CBH II | SDCAA (1) | SDCAA (2) | SDCAA Mean | YPD (1) | YPD (2) | YPD Mean |
|----------------------|------------------|------------------|-------------------|----------------|----------------|-----------------|
| <i>H. jecorina</i> | 19 | 17 | 18 | 0.4 | 0.4 | 0.4 |
| <i>H. jeco C311S</i> | 50 | 43 | 47 | 6.1 | 5.6 | 5.9 |
| <i>H. jeco B7P3</i> | 35 | 33 | 34 | 3.9 | 3.6 | 3.8 |
| <i>H. insolens</i> | 73 | 83 | 78 | 6.2 | 6.0 | 6.1 |
| <i>H. inso C314S</i> | 100 | 97 | 98 | 8.8 | 8.0 | 8.4 |
| <i>H. inso B7P3</i> | 39 | 42 | 40 | 4.4 | 4.1 | 4.2 |

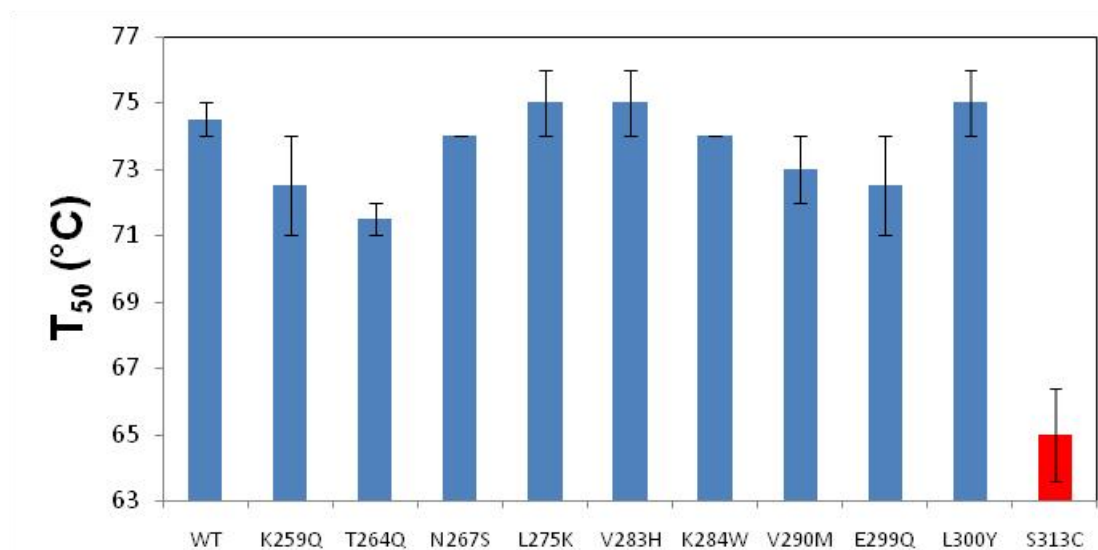
SUPPLEMENTAL FIGURE 1

ClustalW multiple sequence alignment for block 7 from parent 1, *H. insolens* and parent 3, *C.thermophilum*. Arrows denote residues changed in reversion mutants.



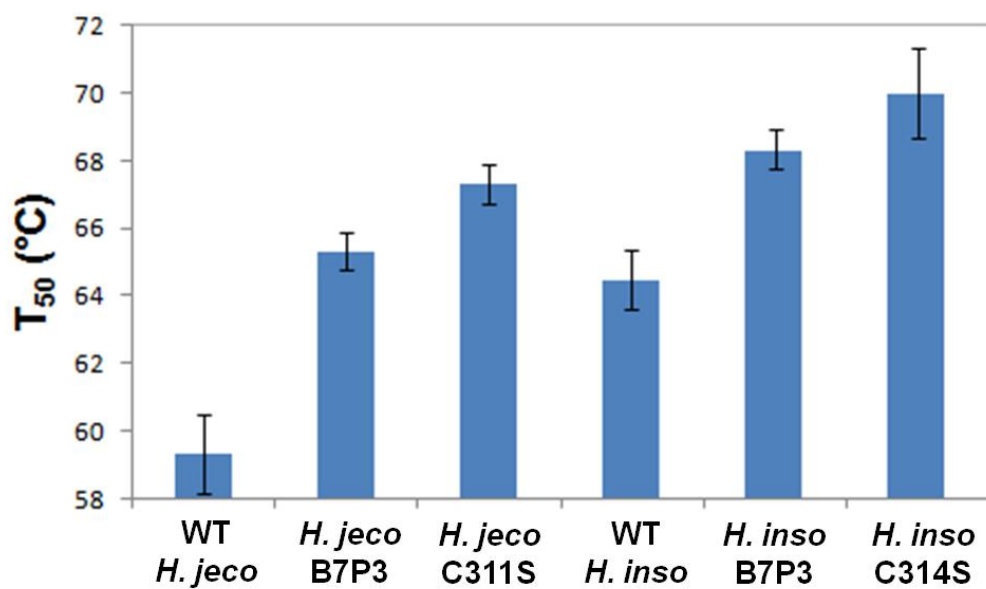
SUPPLEMENTAL FIGURE 2

T_{50} values for 21111331 chimera point mutants. Values shown as average of two independent duplicates, error bars indicate duplicate T_{50} values for each point mutant. Inactivation was carried out for 10 minutes at the temperature being tested in 50 mM sodium acetate buffer, pH 4.8. Residual activity was determined by incubation with 1 g/L phosphoric acid swollen cellulose (PASC) in above buffer for 100 minutes at 50 °C.



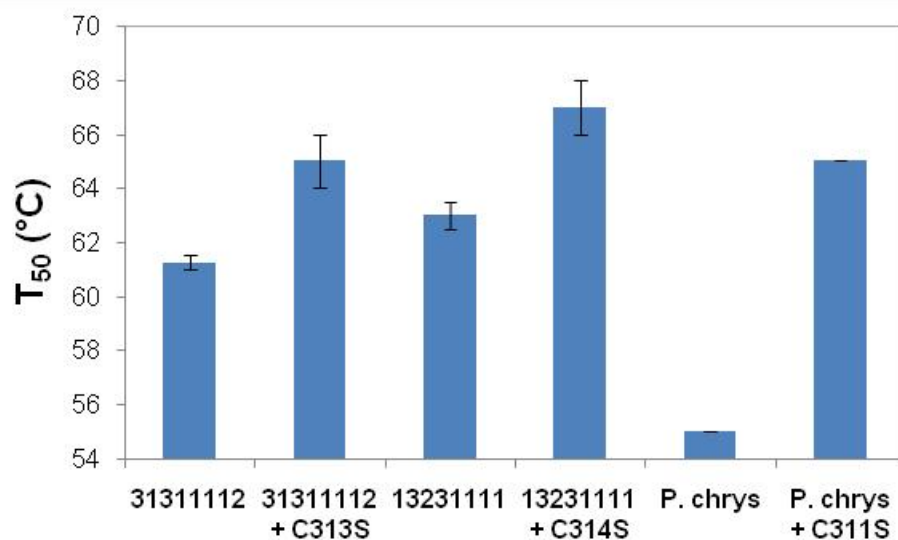
SUPPLEMENTAL FIGURE 3

T_{50} values for *H. insolens* and *H. jecorina* parent CBH IIs, Ser single point mutants and B7P3 block substitution chimeras. Values shown as average of three independent replicates, error bars indicate one standard deviation for each CBH II. Inactivation was carried out for 10 minutes at the temperature being tested in 50 mM sodium acetate buffer, pH 4.8. Residual activity was determined by incubation with 1 g/L phosphoric acid swollen cellulose (PASC) in above buffer for 100 minutes at 50 °C.



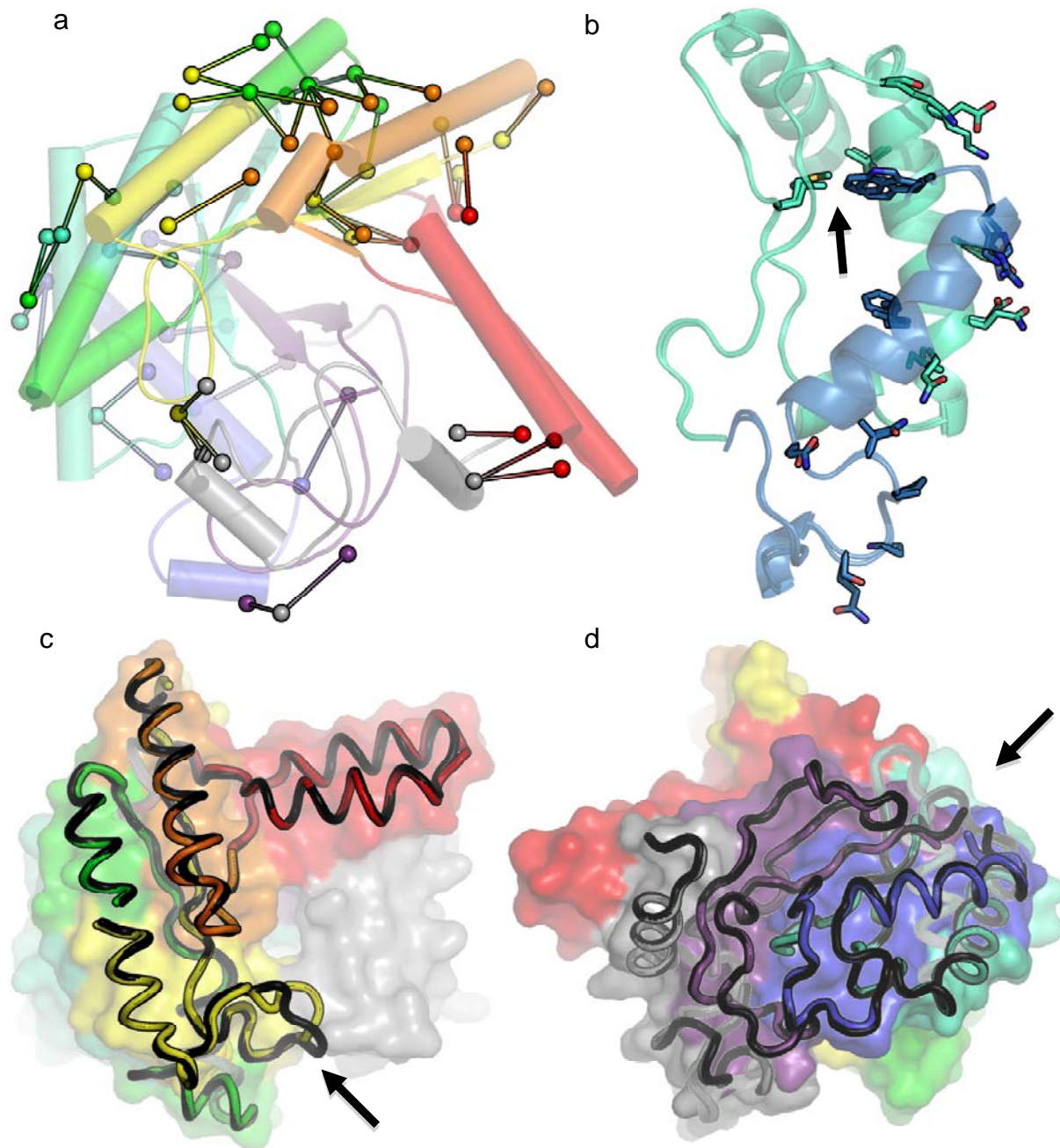
SUPPLEMENTAL FIGURE 4

T_{50} values for CBH II chimeras 31311112, 13231111 and the wild type CBH II catalytic domain from *P. chrysosporium* (fused to the *H. jecorina* CBM) and heterologously secreted from *S. cerevisiae*. Values shown as two independent replicates with error bars indicating values for each trial. Inactivation was carried out for 10 minutes at the temperature tested, in 50 mM sodium acetate buffer, pH 4.8. Residual activity was determined by incubation with 1 g/L phosphoric acid swollen cellulose (PASC) in above buffer for 100 minutes at 50 °C.



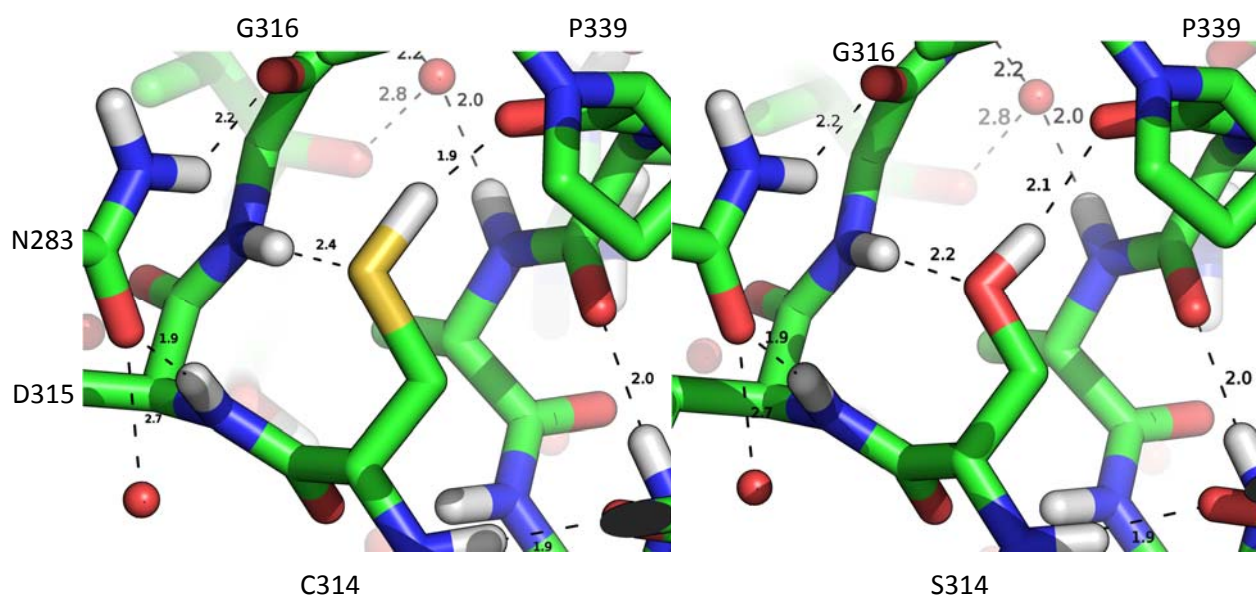
SUPPLEMENTAL FIGURE 5

CBH II recombination block interfaces. (a) Inter-block sites where novel non-parental residue pairs are possible (connected spheres) are usually surface-exposed, potentially allowing solvent to screen the interactions. (b) An example interface (B5-B6) illustrates conservation of the backbone (cartoons for aligned *H. jecorina* and *H. insolens*), variable residues on the surface, and the comparatively rare possibility of a novel buried hydrophobic pair at residues 173 and 253 (arrow). (c) Blocks 1-4 from *H. jecorina* (black cartoon) match cognate *H. insolens* blocks (color-coded cartoon) without large deviations, though movement associated with substrate binding is observed (arrow) in part of B3 (yellow). (d) Cognate blocks 5-8 are also similar, though the indel at the B6,B7 junction (arrow) will require conformational change.



SUPPLEMENTAL FIGURE 6

Structural analysis of C314S mutation and its stabilizing effect. (a) Hydrogen positions for high-resolution *H.insolens* structure (1ocn) were added with REDUCE.¹ (b) The reconfigured geometry of the analogous serine structure was modeled in PyMOL (<http://www.pymol.org>). Sidechain optimization in the SHARPEN² modeling platform (with an all-atom Rosetta energy function) also suggested that both the Cys314 and Ser314 would donate hydrogen bonds to the carbonyl of Pro339, and accept hydrogen bonds from the amide of Gly316. The superior hydrogen bonding capacity of serine may play a role in the greater stability of the serine containing variants. Another possible explanation is geometric complementarity. Specifically, the Cys position from 1ocn shows evidence of conformational strain in that the sidechain is noticeably bent (i.e. the improper dihedral angle from N-C-C α -C β is 6° from the standard position), increasing the distance from the Pro carbonyl. Numbers in figure not preceded by letters denote hydrogen bond distances (Å).



SUPPLEMENTAL FIGURE 7

Alignment of 60 amino acid frame bracketing *H. jecorina* CBH II Cys311 for 196 protein sequences having highest identity to *H. jecorina* CBH II. Residues at 311 equivalent position denoted by shaded bar. Sequences for recombinant *C. thermophilum*, *H. insolens* and *P. chrys* CBH IIs studied in this work are denoted as *C. ther*, *H. inso* and *P. chrys*. Fifty-four of the 250 most identical sequences were excluded due to redundancy (i.e. point mutants for structural studies or >95% identical isoforms).

| | | |
|-----------------|--|-----------------|
| <i>H. jeco</i> | ---T---G---D---S---L---LDSFVWVKPGGEC | ---T---S----- |
| XP_001395308 | ---T---G---D---A---L---VDAFVWVKPGGES | ---T---S----- |
| CAP93233 | ---T---G---D---A---L---ADAFVWVKPGGES | ---T---S----- |
| XP_662886 | ---T---G---D---S---L---LDAFVWVKPGGES | ---T---S----- |
| XP_001552807 | ---T---G---D---A---L---EDAFVWVKPGGES | ---T---S----- |
| XP_001216114 | ---T---G---N---S---L---VDAFVWVKPGGES | ---T---S----- |
| XP_001258843 | ---T---G---D---P---L---QDAFVWVKPGGES | ---T---S----- |
| XP_001392295 | ---T---G---D---E---L---ADAFVWVKPGGES | ---T---S----- |
| XP_002380541 | ---T---G---D---E---L---VDAFVWVKPGGES | ---T---S----- |
| AAQ72468 | ---T---G---D---A---L---VDAFVWVKPGGES | ---T---S----- |
| P46236 | ---T---G---D---A---L---ADAFVWVKPGGES | ---T---S----- |
| XP_001903170 | ---T---G---L---D---I---EDAFVWVKPGGEC | ---T---S----- |
| JC7931 | ---T---G---N---P---L---EDAFVWVKPGGES | ---T---S----- |
| XP_001273717 | ---T---G---D---D---L---LDAFVWVKPGAES | ---T---S----- |
| XP_001210279 | ---T---G---D---E---L---VDAFVWVKPGGES | ---T---S----- |
| XP_002484367 | ---T---G---D---P---L---EDAFVWVKPGGES | ---T---S----- |
| XP_001264772 | ---T---G---D---K---L---VDAFVWVKPGGES | ---T---S----- |
| XP_960770 | ---T---G---S---S---L---ADAFVWVKPGGES | ---T---S----- |
| XP_658877 | ---T---G---D---E---L---VDAFVWVKPGGES | ---T---S----- |
| ACH91035 | ---T---G---D---P---L---EDAFVWVKPGGES | ---T---S----- |
| XP_001226029 | ---T---G---H---D---L---LDAFVWVKPGGEC | ---T---S----- |
| XP_360146 | ---T---G---S---E---L---ADAFVWVKPGGEC | ---V---S----- |
| <i>C. ther</i> | ---T---G---H---E---L---VDAFVWVKPGGES | ---T---S----- |
| XP_002149891 | ---T---G---D---P---L---EDAFVWVKPGGES | ---T---S----- |
| XP_001269265 | ---T---G---D---E---L---VDAFVWVKPGGES | ---T---S----- |
| <i>H. inso</i> | ---T---G---H---Q---Y---VDAFVWVKPGGEC | ---T---S----- |
| XP_001598803 | ---T---G---D---A---L---EDAFVWVKPGGEADG | ---T---S----- |
| AAM76664 | ---T---G---N---A---L---IDAIVWVKPGGES | ---T---S----- |
| XP_001933777 | ---T---G---N---A---N---VDAFVWVKPGGES | ---T---S----- |
| XP_001796781 | ---T---D---D---P---L---LDAFVWVKPGGES | ---T---S----- |
| AAA50608 | ---T---G---S---S---L---IDAIVWVKPGGEC | ---T---S----- |
| AAK28357 | ---T---G---S---S---L---IDSIVWVKPGGEC | ---T---S----- |
| BAF80327 | ---T---G---S---S---L---IDAIVWVKPGGES | ---T---S----- |
| XP_001792324 | ---T---G---N---S---L---VDALVWVKPGGES | ---T---S----- |
| BAG48183 | ---T---G---S---S---L---IDSIVWVKPGGES | ---T---S----- |
| BAH59082 | ---T---G---S---P---L---IDSIVWVKPGGEC | ---T---S----- |
| AAT64008 | ---T---G---S---S---L---IDAIVWVKPGGEC | ---T---T----- |
| <i>P. chrys</i> | ---T---G---S---Q---F---IDSIVWVKPGGEC | ---T---S----- |
| BAH59083 | ---T---P---S---S---L---IDSIVWVKPGGEADG | ---T---S----- |
| XP_001833045 | ---T---P---S---S---A---IDAIVWVKPGGEADG | ---T---S----- |
| AAF35251 | ---T---G---N---P---L---IDAIVWVKPGGES | ---T---S----- |
| XP_001226566 | ---T---G---A---D---L---ADAFVWVKPGGES | ---T---S----- |
| XP_001903893 | ---T---G---L---E---L---ADAFVWVKPGGES | ---T---S----- |
| XP_002391276 | ---T---G---S---S---L---IDSIVWVKPGGES | -----S----- |
| XP_956581 | ---T---G---H---D---L---ADAFVWVKPGGES | ---T---S----- |
| XP_362054 | ---T---G---S---S---L---TDALLWVKPGGES | ---T---S----- |
| XP_001806560 | ---T---G---N---T---L---VDAFVWVKPGGES | ---T---S----- |
| XP_001931623 | ---W---G---D---C---L---VDSFVWVKPGGES | ---T---S----- |
| AAI15038 | ---T---G---S---S---L---IDAIVWVKPGGES | ---T---S----- |
| AAI01211 | ---P---I---S---S---MDYLDIFYWIKPLGES | ---T---S----- |
| XP_001839612 | ---QRILN---N---T---N---VDAIVWVKPGGES | ---T---S----- |
| AAI92497 | ---S---G---M---D---Y---LDAFYWVKPLGES | ---Y---S----- |
| YP_001618727 | ---P---G---G---A---TN-LDAFVWVKPPGDS | ---V---G----- |
| AAM94167 | ---S---G---M---D---Y---LDAFYWVKPYGES | ---T---S----- |
| XP_001836853 | ---Q---GVLQN---P---N---VDAIVWVKPGGES | ---T---S----- |
| AAI01212 | ---S---G---M---D---Y---LDAFYWIKPLGES | ---T---S----- |
| XP_001841133 | ---L---D---S---E---N---VDAIVWVKPGGES | ---T---S----- |
| XP_001841132 | ---Q---AIV-D---S---AN-VDAIVWVKPGGES | ---T---S----- |
| AAF30749 | ---A---G---M---D---Y---LDAFYWVKPYGES | ---T---S----- |
| XP_001224490 | ---T---N---N---T---V---VDALVWVKPGGES | -----S----- |
| XP_368004 | ---T---E---N---P---H---VDSIVWVKPGGES | -----S----- |
| XP_001903209 | ---T---N---N---T---V---VDAIVWVKPGGES | -----S----- |
| AAI51055 | ---P---D---S---SKP-L-LDAYMWIKTPGEADG | ---S---S----- |
| XP_001226518 | ---T---G---N---N---N---VDAIVWVKPGGES | -----S----- |
| XP_957415 | ---Q---SVLQN---S---N---VDAIVWVKPGGES | -----S----- |
| ABY52798 | ---S---G---Y---P---L---LDAFMWLKTPGEADG | ---S---A----- |
| XP_001795501 | ---T---N---N---T---H---IDSIVWVKPGGES | ---A-----S----- |

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| XP_001903645 | ---T---N---N---P---N---VDAILWVKPGGESDG--T---C----- |
| XP_001934153 | ---Q---N---N---T---N---VDSIVWIKPGGESDG--A----- |
| AAF34679 | ---P---D---A---SMP-L-LDAYMWLKTTPGEADG--S---A----- |
| ABY52797 | ---P---S---K---P---L-LDAYMWIKTPGEADG--S---S----- |
| AAC09228 | ---S---G---M---P---L-LDAYMWLKTTPRESDG--S---S----- |
| Q7SIG5 | ---T---N---N---P---N---VDAIVVWVKPGGESDG----- |
| AAR08200 | ---PNP-G---M---P---L-LDAYMWLKTTPGEADG--S---S----- |
| AAD51054 | ---V---N---M---P---L-LDAYMWLKPPAESDG--S---S----- |
| AAB92679 | ---P---D---P---NMP-L-LDAYVWIKTPGESD----- |
| AAB92678 | ---P---N---P---GSMPL-LDAYMWIKTPGEADG--S---S----- |
| AAQ09256 | ---A---G---M---P---L-LDAYMWLKT--PGES--D---G----- |
| AAD02028 | ---P---A---DL---P---G-LDAYVWFKPPGESDG--S---D----- |
| AAC09066 | ---V---N---M---P---L-LDAYMWLKPGGDSDG----- |
| AAP33843 | ---A---G---M---P---L-LDAYMWLKTTPGESDG--S---S----- |
| AA047726 | ---P---D---P---EKYPL-LDAYFWLKPPAESDG--S---S----- |
| AAQ09258 | ---P---A---NY---P---L-LDAFVWIKTPGESDG--N---A----- |
| ZP_01907667 | ---T---A---D---P---R-VDAYLWIKVPGESDG--E---C----- |
| NP_627067 | ---T---A---D---P---L-VDAYLWVKRPGESDG----- |
| ABY52799 | ---S---P---DPEKFP--L-LDAYFWLKPPGEADG--S---D----- |
| YP_879613 | ---T---A---G---A---H-ADAYLWIKRPGESDG--S----- |
| NP_959214 | ---T---A---G---A---H-ADAYLWIKRPGESDG--S----- |
| YP_002186959 | ---T---A---D---P---L-ADAYLWVKRPGESDG--T---C----- |
| YP_001848433 | ---T---A---G---A---H-ADAYLWIKRPGESDG--S----- |
| YP_001132274 | ---T---G---N---P---M-VDAFLWVKRPGESDG----- |
| YP_890960 | ---T---A---G---A---H-ADAYLWVKRPGESDG--A---C----- |
| ZP_04688022 | ---T---A---D---P---L-VDAYLWVKRPGESDG----- |
| YP_642458 | ---T---G---H---P---Q-VDAFLWVKRPGESDG--S---C----- |
| YP_001468205 | ---D---D---D---S---H-QDARLWIKRVGESDG--T----- |
| ZP_04366710 | ---N---D---G---S---G-LDALLWIKLPGESDG----- |
| YP_001506001 | ---T---G---D---S---D-VDAFFWVKIPGESDG--T----- |
| AAD41097 | FIPHR---R---H---C---L-GQTRRRVNPAAKAAG----- |
| ZP_03911849 | ---N---D---S---T---G-LDALLWIKLPGESDG--S---C----- |
| ZP_04475891 | ---T---G---H---D---R-VDAFLWIKRPGESDG----- |
| ZP_04368446 | ---N---D---G---S---G-LDALLWIKRPGESDG--T---C----- |
| ZP_03888455 | ---T---G---Q---P---R-VDAFLWVKRPGESDG--T---C----- |
| ZP_04704893 | ---T---G---D---P---L-LDAYLWIKRPGESDG--S----- |
| NP_334477 | ---T---A---G---A---H-ADAYLWIKRPGESDG--T----- |
| YP_924688 | ---E---G---D---A---EH-LDAYVWVKPPGESDG--E---C----- |
| ZP_03867905 | ---A---S---G---A---H-RGNL-WIKLPGESDG--S---C----- |
| ZP_04747451 | ---T---A---G---A---H-ADAYLWVKRPGESDG--S---C----- |
| YP_002196808 | ---T---G---D---P---L-VDAYLWVKRPGESDG--D----- |
| NP_826394 | ---T---A---D---P---L-VDAYVWVKRPGESDG--T----- |
| YP_001511422 | ---T---G---S---P---H-ADAFWLWIKTPGASDG----- |
| YP_956608 | ---T---G---N---P---M-VDAFLWVKRPGESDG----- |
| NP_778753 | ---P---G---G---H---VDAFQWIKPPPGYSDG--S---SSLIP----- |
| AAC60491 | ---T---G---D---A---N-IDAYLWVKPPGEADG----- |
| NP_298556 | ---P---G---G---H---VDAFQWIKPPPGYSDG--S---SSLIP----- |
| YP_715139 | ---T---G---Y---P---H-ADAFWLWVKTPGASDG----- |
| NP_522144 | ---P---G---G---H---LDAFVWIKPAGEADG--S---SRLIP----- |
| YP_002208806 | ---T---A---D---P---L-VDAYLWVKRPGESDG----- |
| ZP_02961057 | ---D---I---S---A-TVKTAWIKPPGESDG----- |
| YP_001912004 | ---P---G---G---H---IHAYVWGKGGGESDG--S---SKYIP----- |
| NP_638880 | ---P---G---G---H---IHALVWGKGPGESDG--A---S----- |
| AAC09068 | ---V---K---M---P---L-LDAYMWLKTTPGEADG--S---D----- |
| ZP_04704563 | ---T---G---D---D---L-VDAYLWIKRPGESDG----- |
| YP_001705499 | ---T---A---S---P---N-ADAYLWVKRPGESDG--S---C----- |
| P07984 | ---N---D---G---S---G-LDALLWVKLPGESDG----- |
| NP_821732 | ---S---P---D---S---H-LDAFLWIKPPGESDG--A---SSDIP----- |
| ZP_00651218 | ---P---G---G---H---VDAFQWIKPPPGYSDG--S---SSMIP----- |
| YP_872374 | ---P---T---DFPN-A---H-LDAYVWIKPPGESDG--T---S----- |
| YP_431432 | ---PFG-A---G---A---P-VDAFVWIKPPGESDG--T---S----- |
| AAF42880 | ---T---G---D---P---A-IDAYLWIKRPGESDG--T---C----- |
| ZP_04371095 | ---T---G---D---A---A-VDAFLWIKPPGEADG--C---A----- |
| YP_001544904 | ---A---P---V---S---G-IDAYVWVKPPGESDG--V---A----- |
| YP_715138 | ---T---G---N---P---R-ADAYLWIKVPGESDG--A---C----- |
| YP_001826214 | ---T---G---D---E---L-VDAYLWVKRPGESDG--D----- |
| ZP_04696280 | ---T---G---D---E---R-LDAYLWIKRPDGDSDG--T---C----- |
| P50401 | ---P---S---GYAA-S---H-LDAFVWIKPPGESDG--A---STDIP----- |
| ZP_03818362 | ---T---G---D---S---Q-IDAFWLWIKIVGEADG----- |
| NP_625732 | ---T---G---M---G---R-VDAYLWVKLPGESDG----- |
| ZP_00944606 | ---P---G---G---H---LDAVLWIKPPGESDG--S---SRLIP----- |
| ZP_03817628 | ---T---G---D---P---Q-IDAFWLWIKIPGEADG----- |
| ZP_04475504 | ---P---A---A---G-LDAYVWIKPPGESDG--A---SKLIP----- |
| ZP_04331392 | ---T---G---N---P---L-IDAFIWTKLPGESDG----- |
| 2BOE-X | ---T---G---D---P---M-IDAFWLWIKLPGEADG----- |
| YP_001828440 | ---A---P---E---P---G-IDAYVWAKPPGESDG--S---S----- |
| ACR14000 | ---P---A---P---G---VDAFVWVKPQGESDG--I---S----- |
| YP_001823637 | ---T---G---D---D---R-LDAYLWIKRPDGDSDG--T---C----- |
| ZP_01462143 | ---P---A---GYTA-S---R-LDAFVWIKPPGESDG--A---SKEIP----- |
| ZP_04367954 | ---P---A---GAPSAS--H-LDAYVWIKPPGESDG--A---SKEIP----- |
| ZP_04376648 | ---S---G---D---H---V---I-ADVWIKPPGESDG--D---Y----- |

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| YP_001902424 | ----P---G-----A---H--LDAYAWVKPPGESDG--S----STLIP----- |
| ZP_03868469 | ----P---T---DFPN-S---H--LDAYVWVKPPGESDG--S----SSEIP----- |
| ZP_04608509 | ----T---G---D---S---A--IAAYLWVKLPGEADG----- |
| YP_002208593 | ----P---A---T---G-----VDAYVWVKPPGESDG--N----SAAVP----- |
| ZP_03912473 | ----S---G---Y---P---H--IDAFVWVKPPGESDG--S----STEIP----- |
| ZP_04704085 | ----A---P---E---P---G--IDAYVWAKPPGESDG--S----S----- |
| ZP_04707051 | ----T---G---E---A---R--IDAYLWVKLPGESDG--C---S----- |
| ZP_04482497 | ----A---P---E---S---G--IDAYVWVKPPGESDG--S----SKEIP----- |
| BAB83928 | ----P---A-----S---G--IDAYTWIKPPGESDG--N----SAPVD----- |
| ZP_04684292 | ----G---G---A---E---M--L---LWIKVPGESDG--NCGVGA----- |
| YP_002196127 | ----P---P---E---A---G--IDAYLWVKPPGESDG--S----SSEIP----- |
| P33682 | ----G---G---A---E---M--L---LWIKTPGESDG--NCGVGS----- |
| NP_823029 | ----P---A-----T---G--IDAYVWVKPPGESDG--S----S----- |
| NP_828072 | ----T---G---E---A---G--IDAYLWVKLPGESDG----- |
| P26414 | ----T---G---D---P---A--IDAFWLWIKPPGEADG----- |
| YP_925799 | ----V---F---D---G---A--FDGTLWVKHPGESDG--T---K----- |
| ZP_04370960 | ----G---A---S---N---H--IIAYVWIKPPGESDG--D---Y----- |
| ZP_04378286 | ----T---S---A---P---H--ADAYLWIKHPGLSDG--T----- |
| ACR12723 | ----P---S---P---G-----IDAYVWVKPQGESDG--V---S----- |
| ZP_04333277 | ----A---P---E---A---G--IDAYVWVKPPGESDG--S----S----- |
| ZP_04483608 | ----V---G---E---G---A--QDARIWIKVPGESDG----- |
| ZP_03390825 | ----T---T---G---A---H--ADAYLWINGGGGSDG--E---C----- |
| ZP_04377655 | ----T---T---G---A---H--ADAYLWINGGGGSDG--E---C----- |
| YP_001108158 | ----G---G---A---D---L--L---VWAKVPGESDG----- |
| ZP_04377654 | ----T---T---G---A---H--ADAYLWINGGGGSDG--E---C----- |
| ZP_04058051 | ----T---T---G---A---H--ADAYLWINGGGGSDG--E---C----- |
| YP_002203140 | ----P---A-----A---G--IDAYVWIKPPGESDG--A----SSAVP----- |
| YP_632998 | ----V---G---G---A---E--MT--LWIKVPGESDG----- |
| ZP_04684732 | ----A---P---E---P---G--IDAYVWVKPPGESDG--S----SKAID----- |
| ZP_04605447 | ----A---P---E---P---G--IDAYVWVKPPGESDG--S----SKEIP----- |
| YP_288681 | ----P---A---P---G-----VDAYVWVKPPGESDG--A---S----- |
| AAF75786 | ----P---A---P---G-----VDAYVWVKPPGESDG--A---S----- |
| YP_001538764 | ----A---P---E---P---G--IDAYVWVKPPGESDG--S----S----- |
| YP_527744 | ----A---P---E---P---G--VDAYVWVKPQGESDG--I---SDPNFPIDPNDPAK |
| NP_630629 | ----P---A-----A---G--IDAYVWVKPPGESDG--S----SKLID----- |
| ZP_04032911 | ----T---G---H---R---L--VDAFLWVKRPGESDGQCT---R----- |
| YP_001160411 | ----A---P---E---P---G--IDAYVWVKPPGESDG--S----S----- |
| YP_001982934 | ----P---A-----A---G--IDAYVWVKPPGESDG--V---G----- |
| ZP_03817603 | ----P---K---P---N-----IDAYVWIKPPGESDG--S----STQIP----- |
| XP_002396053 | ----T---G---S---S---L--IDAIVWAKPPGESDG--T---S----- |
| ZP_04371165 | ----T---G---V---P---L--VDADLFIKTIGESDGSCCT---R----- |
| AAD54679 | ----V---D---D---P---L--VAGYLWIKVPGESDG--Q---CY----- |
| ZP_03911156 | ----T---G---D---P---L--VAAYLWIKVPGESDG--L----- |
| YP_871896 | ----T---G---V---P---L--LDAYLWVKIPGESDG--Q---C----- |